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October 9, 2008

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Re: Title: MODULATION OF HIF1ALPHA AND HIF2ALPHA EXPRESSION  
Letters Patent No. 7,217,572  
Issued: May 15, 2007  
Our Reference: ISIS.038CP1

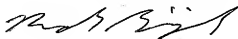
Dear Sir:

Enclosed for filing is a Certificate of Correction in connection with the above-identified patent.

As some of the errors cited in the Certificate of Correction were incurred through the fault of the Applicant, the fee of \$100 will be paid via EFS Web. Please charge any additional fees to our Deposit Account No. 11-1410.

Respectfully submitted,

Knobbe, Martens, Olson & Bear, LLP



Brenden Gingrich  
Registration No. 60,295  
Customer No. 55,389

Enclosures

5927326  
091508

Orange County  
949-760-0404

San Francisco  
415-954-4114

Los Angeles  
310-551-3450

Riverside  
951-781-9231

Seattle  
206-405-2000

Washington, DC  
202-640-6400

# UNITED STATES PATENT AND TRADEMARK OFFICE

## CERTIFICATE OF CORRECTION

**PATENT NO.** : 7,217,572  
**APPLICATION NO.** : 10/719,370  
**ISSUE DATE** : May 15, 2007  
**INVENTOR(S)** : Ward et al.

Page 1 of 3

It is certified that error appears in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

First Page, Col. 2 (Other Publications), line 6, Delete "pates" and insert -- pages --, therefor.  
First Page, Col. 2 (Other Publications), line 9, Delete "inhibitor-1<sup>'''</sup>" and insert -- inhibitor-1, --, therefor.  
Page 2, Col. 1 (Other Publications), line 14, Delete "hypoxia inducible" and insert -- hypoxia-inducible --, therefor.  
Page 2, Col. 1 (Other Publications), line 34, Delete "Hypoxia Inducible" and insert -- Hypoxia-Inducible --, therefor.  
Page 2, Col. 1 (Other Publications), line 57, Delete "hypoxia inducible" and insert -- hypoxia-inducible --, therefor.  
Page 2, Col. 1 (Other Publications), line 61, Delete "Hypoxia inducible" and insert -- Hypoxia-inducible --, therefor.  
Page 2, Col. 2 (Other Publications), line 1, Delete "s" and insert -- a --, therefor.  
Page 2, Col. 2 (Other Publications), line 72, Delete "Endotelial" and insert -- Endothelial --, therefor.  
Page 3, Col. 2 (Other Publications), line 6, Delete "Investigation," and insert -- Investigation, --, therefor.  
Col. 1, line 52, Delete "hypoxia inducible" and insert -- hypoxia-inducible --, therefor.  
Col. 5, line 13 (Approx.), Delete "bindingsite" and insert -- binding site --, therefor. (Consider Space)  
Col. 5, line 57, Delete "(PPARS)" and insert -- (PPARs) --, therefor.  
Col. 7, line 51 (Approx.), Delete "HIF2aactivity" and insert -- HIF2a activity --, therefor. (Consider Space)  
Col. 23, line 51 (Approx.), Delete "B.ed," and insert -- B.,ed., --, therefor.  
Col. 26, line 39, Delete "administration," and insert -- administration --, therefor.  
Col. 28, line 63, Delete "ara-binoside," and insert -- arabinoside, --, therefor.  
Col. 29, line 3, Delete "deoxycyco-formycin," and insert -- deoxycycoformycin, --, therefor.  
Col. 29, line 8, Delete "teni-poside," and insert -- teniposide, --, therefor.  
Col. 33, line 15, Delete "Tetrahedrom" and insert -- Tetrahedron --, therefor.  
Col. 34, line 62 (Approx.), After "Strand" insert -- (SEQ ID NO: 456) --.  
Col. 34, line 63 (Approx.), After "Complement" insert -- (SEQ ID NO: 457) --.  
Col. 35, line 5, After "Strand" insert -- (SEQ ID NO: 455) --.  
Col. 35, line 6 (Approx.), After "Complement" insert -- (SEQ ID NO: 458) --.  
Col. 35, line 25, Delete "HIF1● or HIF2●" and insert -- HIF1α or HIF2α --, therefor.  
Col. 35, line 29, Delete "200 ●L" and insert -- 200 μL --, therefor.  
Col. 35, line 31, Delete "130 ●L" and insert -- 130 μL --, therefor.  
Col. 35, line 32, Delete "●g/mL" and insert -- μg/mL --, therefor.  
Col. 35, line 33, Delete "200 M." and insert -- 200 nM. --, therefor.  
Col. 39, line 61 (Approx.), Delete "minutes," and insert -- minutes. --, therefor.  
Col. 59, line 22, Col. 4 (Table 4), Delete "tccaatggtgacaactgac" and insert -- tccaatggtgacaactgac --, therefor.  
Col. 62, lines 48-49, Delete "AGGCCGAGAATGGGAAGCTTGTCATC" and insert --  
AAGGCCGAGAATGGGAAGCTTGTCATC --, therefor.  
Col. 66, line 57, Delete "3'-directions)" and insert -- 3' directions) --, therefor.  
Col. 79, line 4, Delete "Hif1α" and insert -- HIF1α --, therefor.

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DOCKET NO. ISIS.038CP1

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Page 2 of 3

Col. 79, line 11 (Approx.), Delete "Hif2a" and insert -- HIF2a --, therefor.  
Col. 82, lines 34-35, Delete "HIF1● and not HIF2●" and insert -- HIF1a and not HIF2a --, therefor.  
Col. 82, line 37, Delete "HIF2● and not HIF1●" and insert -- HIF2a and not HIF1a --, therefor.  
Col. 82, line 39, Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 82, line 40, Delete "HIF2●" and insert -- HIF2a --, therefor.  
Col. 82, line 62 (Approx.), Delete "(TCGCGGCTGGATTAC;)" and insert -- (TCGCGGCTGGACGATTAC; --, therefor.  
Col. 82, line 64, Delete "HIF1● and HIF2●" and insert -- HIF1a and HIF2a --, therefor.  
Col. 82, line 66, Delete "HIF1● and HIF2●" and insert -- HIF1a and HIF2a --, therefor.  
Col. 83-84, lines 2-3, Col. 6 (Table 13), Delete "Mismatches" and insert -- Mismatch --, therefor.  
Col. 84, line 11 (Approx.), Delete "HIF1●/HIF2●" and insert -- HIF1a/HIF2a --, therefor.  
Col. 85, line 8 (Approx.), Delete "Crossr Acting" and insert -- Crossreacting --, therefor.  
Col. 87, lines 1-5 (Approx.), Delete "Table 15 Effect of HIF1a and HIF2a antisense oligonucleotides on angiogenic tube formation".  
Col. 87, line 20, Insert -- Table 15 Effect of HIF1a and HIF2a antisense oligonucleotides on angiogenic tube formation -  
Col. 87, line 53, Delete "HIF1● and crossreactive to human HIF1●" and insert -- HIF1a and crossreactive to human HIF1a --, therefor.  
Col. 87, lines 60-61 (Approx.), Delete "HIF1● expression in mouse liver by antisense to HIF1●" and insert -- HIF1a expression in mouse liver by antisense to HIF1a --, therefor.  
Col. 87, line 62 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, lines 3-4 (Approx.), Delete "HIF1● expression in mouse liver by antisense to HIF1●" and insert -- HIF1a expression in mouse liver by antisense to HIF1a --, therefor.  
Col. 88, line 6 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, line 17 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, line 31 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, line 54, Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, line 61 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 88, line 63 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 89, line 4 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 89, line 5 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 89, line 11 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 89, line 17 (Approx.), Delete "HIF1●" and insert -- HIF1a --, therefor.  
Col. 89, line 42, Delete "HIF2●" and insert -- HIF2a --, therefor.  
Col. 89, line 43, Delete "HIF2●" and insert -- HIF2a --, therefor.  
Col. 89, line 46, Delete "HIF2●" and insert -- HIF2a --, therefor.  
Col. 89, line 47, Delete "HIF2●" and insert -- HIF2a --, therefor.

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Page 3 of 3

Col. 92, line 11, Delete "10 •M" and insert -- 10  $\mu$ M --, therefor.

In the Claims:

Col. 366, line 18 (Approx.), In Claim 17, delete "oligonacleotide" and insert -- oligonucleotide --, therefor.

Col. 366, line 23 (Approx.), In Claim 17, delete "5-niethylcytidines." and insert -- 5-methylcytidines. --, therefor.

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